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(54) Title: SECRETED FACTORS

(57) Abstract: The invention concerns new secreted factors encoded by clones P00184 D11 (SEQ ID NO:1), P00185 D11(SEQ ID NO:3), P00188 D12 (SEQ ID NO:5), P00188 E01 (SEQ ID NO:7), P00194 G01 (SEQ ID NO:9), P00194 G05 (SEQ ID NO:11), P00194 H10 (SEQ ID NO:13), P00199 D08 (SEQ ID NO:15), P00203 D04 (SEQ ID NO:17), P00203 E06 (SEQ ID NO:19), P00209 F06 (SEQ ID NO:21), P00219 D02 (SEQ ID NO:23), P00219 F06 (SEQ ID NO:25), P00220 H05 (SEQ ID NO:27), P00222 G03 (SEQ ID NO:29), P00225 C01 (SEQ ID NO:32), P00227 D11 (SEQ ID NO:34), P00228 F03 (SEQ ID NO:36), P00233 H08 (SEQ ID NO:38), P00235 G08 (SEQ ID NO:40), P00239 C11 (SEQ ID NO:42), P00240 E05 (SEQ ID NO:45), P00247 A04 (SEQ ID NO:50), P00248 B04 (SEQ ID NO:52), P00249 F09 (SEQ ID NO:54), P00258 A10 (SEQ ID NO:56), P00262 C10 (SEQ ID NO:58), P00269 H08 (SEQ ID NO:62), P00628 H02 (SEQ ID NO:66), P00629 C08 (SEQ ID NO:68), P00641 G11 (SEQ ID NO:71), P00648 E12 (SEQ ID NO:73), P00697 C03 (SEQ ID NO:75), and other mammalian homologues and variants of such factor, as well as polynucleotides encoding them. The invention further concerns methods and means for producing such factors and their use in the diagnosis and treatment of various cardiac, renal or inflammatory diseases.

Seq ID	ClassID	PKD Area	Kidney		Heart									
			No.	Hyp	2w	4w	8w	12w	16w	2w	4w	8w		
135	P00001	PKD	—	—	▲	▲	▲	▲	▲	—	▲	—	▲	▲
136	P00002	PKD	—	—	▲	▲	▲	▲	▲	—	—	—	—	—
137	P00003	PKD	—	▲	—	▼	—	—	—	—	—	—	—	—
138	P00004	PKD	—	▲	—	—	—	—	—	—	—	—	—	—
139	P00005	PKD	—	—	▼	—	▲	—	▲	—	—	—	—	—
140	P00006	PKD	—	▲	—	▼	—	▼	▼	—	—	—	—	—
141	P00012_C04	PKD	—	—	—	—	▼	—	—	—	—	▼	—	—
142	P00012_H04	PKD	—	—	—	▼	▼	▼	—	—	▼	▼	▼	▼
143	P00012_H02	PKD	▲	—	—	—	—	▲	▼	▼	—	—	—	—
144	P00012_C01	PKD	—	—	—	—	▼	—	—	—	—	—	—	—
145	P00012_S01	PKD	—	—	—	—	—	—	—	—	—	—	—	—
146	P00012_G01	PKD	—	—	▲	—	—	—	—	—	—	—	—	—
147	P00012_G02	PKD	—	—	—	—	—	—	—	—	—	—	—	—
148	P00027_C03	PKD	▼	—	—	—	—	—	—	▼	▼	—	—	—

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patent (AM, AZ, BY, KG, KZ, MD, RU, TJ, TM), European patent (AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR), OAPI patent (BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG).

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CLAIMS:

1. An isolated nucleic acid molecule comprising a poly- or oligonucleotide selected from the group consisting of:
 - (a) a polynucleotide encoding a polypeptide having at least about 80% sequence identity with any amino acid sequence selected from the group consisting of: amino acids 1 to 193 of SEQ ID NO: 4, amino acids 1 to 236 of SEQ ID NO:6, amino acids 1 to 61 of SEQ ID NO: 8, amino acids 1 to 92 of SEQ ID NO:12, amino acids 1 to 86 of SEQ ID NO:14, amino acids 1 to 36 of SEQ ID NO:16, amino acids 1 to 83 of SEQ ID NO:18, amino acids 1 to 82 of SEQ ID NO:20, amino acids 1 to 462 of SEQ ID NO:22, amino acids 1 to 170 of SEQ ID NO:24, amino acids -26 to 233 of Fig. 13 (amino acids 1 to 259 of SEQ ID NO: 26), amino acids 1 to 30 of SEQ ID NO:28, amino acids 1 to 30 of SEQ ID NO:35, amino acids 1 to 100 of SEQ ID NO:37, amino acids 1 to 65 of SEQ ID NO:39, amino acids 1 to 46 of SEQ ID NO:43, amino acids 1 to 313 of SEQ ID NO:46, amino acids 1 to 58 of SEQ ID NO:51, amino acids -35 to 387 of Fig. 29 (amino acids 1 to 422 of SEQ ID NO: 53), amino acids 1 to 58 of SEQ ID NO:55, amino acids 1 to 52 of SEQ ID NO:57, amino acids 1 to 245 of SEQ ID NO:59, amino acids 1 to 142 of SEQ ID NO:63, amino acids 1 to 49 of SEQ ID NO:67, amino acids 1 to 70 of SEQ ID NO:69, amino acids 1 to 113 of SEQ ID NO: 72, and amino acids 1 to 97 of SEQ ID NO:76; or a transmembrane domain (membrane spanning segment/region) deleted or inactivated variant thereof;
 - (b) a polynucleotide encoding a polypeptide of amino acids 1 to 233 of SEQ ID NO: 26, or amino acids 1 to 387 of SEQ ID NO: 53;
 - (c) a polynucleotide encoding amino acids 1 to 203 of SEQ ID NO: 2, amino acids 1 to 193 of SEQ ID NO: 4, amino acids 1 to 236 of SEQ ID NO:6, amino acids 1 to 61 of SEQ ID NO: 8, amino acids 1 to 79 of SEQ ID NO:10, amino acids 1 to 92 of SEQ ID NO:12, amino acids 1 to 86 of SEQ ID NO:14, amino acids 1 to 36 of SEQ ID NO:16, amino acids 1 to 83 of SEQ ID NO:18, amino acids 1 to 82 of SEQ ID NO:20, amino acids 1 to 462 of SEQ ID NO:22, amino acids 1 to 170 of SEQ ID NO:24, amino acids -26 to 233 of Fig. 13 (amino acids 1 to 259 of SEQ ID NO:26), amino acids 1 to 30 of SEQ ID NO:28, amino acids 1 to 39 of SEQ ID NO:30, amino acids 1 to 541 of SEQ ID NO: 33, amino acids 1 to 30 of SEQ ID NO:35, amino acids 1 to 100 of SEQ ID NO:37, amino acids 1 to 65 of SEQ ID NO:39, amino acids 1 to 42 of SEQ ID NO:41, amino acids 1 to 46 of SEQ ID NO:43, amino acids 1 to 313 of SEQ ID NO:46, amino acids 1 to 58 of SEQ ID NO:51, amino acids -35 to 387 of Fig. 29 (amino acids 1 to 422 of SEQ ID NO:53), amino acids 1 to 58 of SEQ ID NO:55, amino acids 1 to 52 of SEQ ID NO:57, amino acids 1 to 245 of SEQ ID NO:59, amino acids 1 to 142 of SEQ ID NO:63, amino acids 1 to 49 of SEQ ID NO:67, amino acids 1 to 70 of SEQ ID NO:69, amino acids 1 to 113 of SEQ ID NO: 72, and amino acids 1 to 114 of SEQ ID NO:74, and amino acids 1 to 97 of SEQ ID NO:76; or a transmembrane domain (membrane spanning segment/region) deleted or inactivated variant thereof.
 - (d) a polynucleotide hybridizing under stringent conditions with the complement of the coding region of SEQ ID NO: 1, wherein said polynucleotide encodes a polypeptide having at least one biological activity of the polypeptide encoded by clone P00184_D11 (SEQ ID NO: 1), a polynucleotide hybridizing

under stringent conditions with the complement of the coding region of SEQ ID NO: 3, wherein said polynucleotide encodes a polypeptide having at least one biological activity of the polypeptide encoded by clone P00185_D11 (SEQ ID NO: 3); a polynucleotide hybridizing under stringent conditions with the complement of the coding region of SEQ ID NO: 5, wherein said polynucleotide encodes a polypeptide having at least one biological activity of the polypeptide encoded by clone P00188_D12 (SEQ ID NO: 5), a polynucleotide hybridizing under stringent conditions with the complement of the coding region of SEQ ID NO: 7, wherein said polynucleotide encodes a polypeptide having at least one biological activity of the polypeptide encoded by clone P00188_E01 (SEQ ID NO: 7), a polynucleotide hybridizing under stringent conditions with the complement of the coding region of SEQ ID NO: 9, wherein said polynucleotide encodes a polypeptide having at least one biological activity of the polypeptide encoded by clone P00194_G01 (SEQ ID NO: 9), a polynucleotide hybridizing under stringent conditions with the complement of the coding region of SEQ ID NO: 11, wherein said polynucleotide encodes a polypeptide having at least one biological activity of the polypeptide encoded by clone P00194_G05 (SEQ ID NO: 11), a polynucleotide hybridizing under stringent conditions with the complement of the coding region of SEQ ID NO: 13, wherein said polynucleotide encodes a polypeptide having at least one biological activity of the polypeptide encoded by clone P00194_H10 (SEQ ID NO: 13), a polynucleotide hybridizing under stringent conditions with the complement of the coding region of SEQ ID NO: 15, wherein said polynucleotide encodes a polypeptide having at least one biological activity of the polypeptide encoded by clone P00199_D08 (SEQ ID NO: 15), a polynucleotide hybridizing under stringent conditions with the complement of the coding region of SEQ ID NO: 17, wherein said polynucleotide encodes a polypeptide having at least one biological activity of the polypeptide encoded by clone P00203_D04 (SEQ ID NO: 17), a polynucleotide hybridizing under stringent conditions with the complement of the coding region of SEQ ID NO: 19, wherein said polynucleotide encodes a polypeptide having at least one biological activity of the polypeptide encoded by clone P00203_E06 (SEQ ID NO: 19), a polynucleotide hybridizing under stringent conditions with the complement of the coding region of SEQ ID NO: 21, wherein said polynucleotide encodes a polypeptide having at least one biological activity of the polypeptide encoded by clone P00209_F06 (SEQ ID NO: 21), a polynucleotide hybridizing under stringent conditions with the complement of the coding region of SEQ ID NO: 23, wherein said polynucleotide encodes a polypeptide having at least one biological activity of the polypeptide encoded by clone P00219_D02 (SEQ ID NO: 23), a polynucleotide hybridizing under stringent conditions with the complement of the coding region of SEQ ID NO: 25, wherein said polynucleotide encodes a polypeptide having at least one biological activity of the polypeptide encoded by clone P00219_F06 (SEQ ID NO: 25), a polynucleotide hybridizing under stringent conditions with the complement of the coding region of SEQ ID NO: 27, wherein said polynucleotide encodes a polypeptide having at least one biological activity of the polypeptide encoded by clone P00220_H05 (SEQ ID NO: 27), a polynucleotide hybridizing under stringent conditions with the complement of the coding region of SEQ ID NO: 29, wherein said polynucleotide encodes a polypeptide having at least one biological activity of the polypeptide encoded by clone P00222_G03 (SEQ ID NO: 29),

a polynucleotide hybridizing under stringent conditions with the complement of the polynucleotide of SEQ ID NO: 31 (clone P00223_F07), a polynucleotide hybridizing under stringent conditions with the complement of the coding region of SEQ ID NO: 32, wherein said polynucleotide encodes a polypeptide having at least one biological activity of the polypeptide encoded by clone P00225_C01 (SEQ ID NO: 32),
5 a polynucleotide hybridizing under stringent conditions with the complement of the coding region of SEQ ID NO: 34, wherein said polynucleotide encodes a polypeptide having at least one biological activity of the polypeptide encoded by clone P00227_D11 (SEQ ID NO: 34), a polynucleotide hybridizing under stringent conditions with the complement of the coding region of SEQ ID NO: 36, wherein said polynucleotide encodes a polypeptide having at least one biological activity of the polypeptide encoded by
10 clone P00228_F03 (SEQ ID NO: 36), a polynucleotide hybridizing under stringent conditions with the complement of the coding region of SEQ ID NO: 38, wherein said polynucleotide encodes a polypeptide having at least one biological activity of the polypeptide encoded by clone P00233_H08 (SEQ ID NO: 38),
a polynucleotide hybridizing under stringent conditions with the complement of the coding region of SEQ
15 ID NO: 40, wherein said polynucleotide encodes a polypeptide having at least one biological activity of the polypeptide encoded by clone P00235_G08 (SEQ ID NO: 40), a polynucleotide hybridizing under stringent conditions with the complement of the coding region of SEQ ID NO: 42, wherein said polynucleotide encodes a polypeptide having at least one biological activity of the polypeptide encoded by
20 clone P00239_C11 (SEQ ID NO: 42), a polynucleotide hybridizing under stringent conditions with the complement of the polynucleotide of SEQ ID NO: 44 (clone P00240_B04), a polynucleotide hybridizing under stringent conditions with the complement of the coding region of SEQ ID NO: 45, wherein said polynucleotide encodes a polypeptide having at least one biological activity of the polypeptide encoded by
25 clone P00240_E05 (SEQ ID NO: 45), a polynucleotide hybridizing under stringent conditions with the complement of the polynucleotide of SEQ ID NO: 47 (clone P00241_E12), a polynucleotide hybridizing under stringent conditions with the complement of the polynucleotide of SEQ ID NO: 48 (clone P00245_D06), a polynucleotide hybridizing under stringent conditions with the complement of the polynucleotide of SEQ ID NO: 49 (clone P00246_D12), a polynucleotide hybridizing under stringent conditions with the complement of the coding region of SEQ ID NO: 50, wherein said polynucleotide encodes a polypeptide having at least one biological activity of the polypeptide encoded by clone
30 P00247_A04 (SEQ ID NO: 50), a polynucleotide hybridizing under stringent conditions with the complement of the coding region of SEQ ID NO: 52, wherein said polynucleotide encodes a polypeptide having at least one biological activity of the polypeptide encoded by clone P00248_B04 (SEQ ID NO: 52),
a polynucleotide hybridizing under stringent conditions with the complement of the coding region of SEQ
35 ID NO: 54, wherein said polynucleotide encodes a polypeptide having at least one biological activity of the polypeptide encoded by clone P00249_F09 (SEQ ID NO: 54), a polynucleotide hybridizing under stringent conditions with the complement of the coding region of SEQ ID NO: 56, wherein said polynucleotide encodes a polypeptide having at least one biological activity of the polypeptide encoded by clone P00258_A10 (SEQ ID NO: 56), a polynucleotide hybridizing under stringent conditions with the

complement of the coding region of SEQ ID NO: 58, wherein said polynucleotide encodes a polypeptide having at least one biological activity of the polypeptide encoded by clone P00262_C10 (SEQ ID NO: 58), a polynucleotide hybridizing under stringent conditions with the complement of the polynucleotide of SEQ ID NO: 60 (clone P00263_G06), a polynucleotide hybridizing under stringent conditions with the complement of the polynucleotide of SEQ ID NO: 61 (clone P00267_F08), a polynucleotide hybridizing under stringent conditions with the complement of the coding region of SEQ ID NO: 62, wherein said polynucleotide encodes a polypeptide having at least one biological activity of the polypeptide encoded by clone P00269_H08 (SEQ ID NO: 62), a polynucleotide hybridizing under stringent conditions with the complement of the polynucleotide of SEQ ID NO: 64 (clone P00312_C04), a polynucleotide hybridizing under stringent conditions with the complement of the polynucleotide of SEQ ID NO: 65 (clone P00324_H02), a polynucleotide hybridizing under stringent conditions with the complement of the coding region of SEQ ID NO: 66, wherein said polynucleotide encodes a polypeptide having at least one biological activity of the polypeptide encoded by clone P00628_H02 (SEQ ID NO: 66), a polynucleotide hybridizing under stringent conditions with the complement of the coding region of SEQ ID NO: 68, wherein said polynucleotide encodes a polypeptide having at least one biological activity of the polypeptide encoded by clone P00629_C08 (SEQ ID NO: 68), a polynucleotide hybridizing under stringent conditions with the complement of the polynucleotide of SEQ ID NO: 70 (clone P00634_G11), a polynucleotide hybridizing under stringent conditions with the complement of the coding region of SEQ ID NO: 71, wherein said polynucleotide encodes a polypeptide having at least one biological activity of the polypeptide encoded by clone P00641_G11 (SEQ ID NO: 71), a polynucleotide hybridizing under stringent conditions with the complement of the coding region of SEQ ID NO: 73, wherein said polynucleotide encodes a polypeptide having at least one biological activity of the polypeptide encoded by clone P00648_E12 (SEQ ID NO: 73), a polynucleotide hybridizing under stringent conditions with the complement of the coding region of SEQ ID NO: 75, wherein said polynucleotide encodes a polypeptide having at least one biological activity of the polypeptide encoded by clone P00697_C03 (SEQ ID NO: 75);

(e) a polynucleotide encoding at least about 50 contiguous amino acids from amino acids 1 to 148 of SEQ ID NO: 2, wherein said polynucleotide encodes a polypeptide having at least one biological activity of the polypeptide encoded by clone P00184_D11 (SEQ ID NO: 1), a polynucleotide encoding at least about 50 contiguous amino acids from amino acids 1 to 193 of SEQ ID NO: 4, wherein said polynucleotide encodes a polypeptide having at least one biological activity of the polypeptide encoded by clone P00185_D11 (SEQ ID NO: 3); a polynucleotide encoding at least about 50 contiguous amino acids from amino acids 1 to 236 of SEQ ID NO: 6, wherein said polynucleotide encodes a polypeptide having at least one biological activity of the polypeptide encoded by clone P00188_D12 (SEQ ID NO: 5), a polynucleotide encoding at least about 50 contiguous amino acids from amino acids 1 to 61 of SEQ ID NO: 8, wherein said polynucleotide encodes a polypeptide having at least one biological activity of the polypeptide encoded by clone P00188_E01 (SEQ ID NO: 7), a polynucleotide encoding at least about 50 contiguous amino acids from amino acids 1 to 79 of SEQ ID NO: 10, wherein said polynucleotide encodes

a polypeptide having at least one biological activity of the polypeptide encoded by clone P00194_G01 (SEQ ID NO: 9), a polynucleotide encoding at least about 50 contiguous amino acids from amino acids 1 to 92 of SEQ ID NO: 12, wherein said polynucleotide encodes a polypeptide having at least one biological activity of the polypeptide encoded by clone P00194_G05 (SEQ ID NO: 11), a polynucleotide encoding at least about 50 contiguous amino acids from amino acids 1 to 86 of SEQ ID NO: 14, wherein said polynucleotide encodes a polypeptide having at least one biological activity of the polypeptide encoded by clone P00194_H10 (SEQ ID NO: 13), a polynucleotide encoding at least about 50 contiguous amino acids from amino acids 1 to 36 of SEQ ID NO: 16, wherein said polynucleotide encodes a polypeptide having at least one biological activity of the polypeptide encoded by clone P00199_D08 (SEQ ID NO: 15), a polynucleotide encoding at least about 50 contiguous amino acids from amino acids 1 to 83 of SEQ ID NO: 18, wherein said polynucleotide encodes a polypeptide having at least one biological activity of the polypeptide encoded by clone P00203_D04 (SEQ ID NO: 17), a polynucleotide encoding at least about 50 contiguous amino acids from amino acids 1 to 82 of SEQ ID NO: 20, wherein said polynucleotide encodes a polypeptide having at least one biological activity of the polypeptide encoded by clone P00203_E06 (SEQ ID NO: 19), a polynucleotide encoding at least about 50 contiguous amino acids from amino acids 1 to 462 of SEQ ID NO: 22, wherein said polynucleotide encodes a polypeptide having at least one biological activity of the polypeptide encoded by clone P00209_F06 (SEQ ID NO: 21), a polynucleotide encoding at least about 50 contiguous amino acids from amino acids 1 to 170 of SEQ ID NO: 24, wherein said polynucleotide encodes a polypeptide having at least one biological activity of the polypeptide encoded by clone P00219_D02 (SEQ ID NO: 23), a polynucleotide encoding at least about 50 contiguous amino acids from amino acids -26 to 233 of Fig. 13 (amino acids 1 to 259 of SEQ ID NO: 26), wherein said polynucleotide encodes a polypeptide having at least one biological activity of the polypeptide encoded by clone P00219_F06 (SEQ ID NO: 25), a polynucleotide encoding at least about 50 contiguous amino acids from amino acids 1 to 30 of SEQ ID NO: 28, wherein said polynucleotide encodes a polypeptide having at least one biological activity of the polypeptide encoded by clone P00220_H05 (SEQ ID NO: 27), a polynucleotide encoding at least about 50 contiguous amino acids from amino acids 1 to 39 of SEQ ID NO: 30, wherein said polynucleotide encodes a polypeptide having at least one biological activity of the polypeptide encoded by clone P00222_G03 (SEQ ID NO: 29), a polynucleotide encoding at least about 50 contiguous amino acids from amino acids 1 to 541 of SEQ ID NO: 33, wherein said polynucleotide encodes a polypeptide having at least one biological activity of the polypeptide encoded by clone P00225_C01 (SEQ ID NO: 32), a polynucleotide encoding at least about 50 contiguous amino acids from amino acids 1 to 30 of SEQ ID NO: 35, wherein said polynucleotide encodes a polypeptide having at least one biological activity of the polypeptide encoded by clone P00227_D11 (SEQ ID NO: 34), a polynucleotide encoding at least about 50 contiguous amino acids from amino acids 1 to 100 of SEQ ID NO: 37, wherein said polynucleotide encodes a polypeptide having at least one biological activity of the polypeptide encoded by clone P00228_F03 (SEQ ID NO: 36), a polynucleotide encoding at least about 50 contiguous amino acids from amino acids 1 to 65 of SEQ ID NO: 39, wherein said polynucleotide encodes

a polypeptide having at least one biological activity of the polypeptide encoded by clone P00233_H08 (SEQ ID NO: 38), a polynucleotide encoding at least about 50 contiguous amino acids from amino acids 1 to 41 of SEQ ID NO: 39, wherein said polynucleotide encodes a polypeptide having at least one biological activity of the polypeptide encoded by clone P00235_G08 (SEQ ID NO: 40), a polynucleotide encoding at least about 50 contiguous amino acids from amino acids 1 to 46 of SEQ ID NO: 43, wherein said polynucleotide encodes a polypeptide having at least one biological activity of the polypeptide encoded by clone P00239_C11 (SEQ ID NO: 42), a polynucleotide encoding at least about 50 contiguous amino acids from amino acids 1 to 313 of SEQ ID NO: 46, wherein said polynucleotide encodes a polypeptide having at least one biological activity of the polypeptide encoded by clone P00240_E05 (SEQ ID NO: 45), a polynucleotide encoding at least about 50 contiguous amino acids from amino acids 1 to 58 of SEQ ID NO: 51, wherein said polynucleotide encodes a polypeptide having at least one biological activity of the polypeptide encoded by clone P00247_A04 (SEQ ID NO: 50), a polynucleotide encoding at least about 50 contiguous amino acids from amino acids -35 to 387 of Fig. 29 (amino acids 1 to 422 of SEQ ID NO: 53), wherein said polynucleotide encodes a polypeptide having at least one biological activity of the polypeptide encoded by clone P00248_B04 (SEQ ID NO: 52), a polynucleotide encoding at least about 50 contiguous amino acids from amino acids 1 to 58 of SEQ ID NO: 55, wherein said polynucleotide encodes a polypeptide having at least one biological activity of the polypeptide encoded by clone P00249_F09 (SEQ ID NO: 54), a polynucleotide encoding at least about 50 contiguous amino acids from amino acids 1 to 52 of SEQ ID NO: 57, wherein said polynucleotide encodes a polypeptide having at least one biological activity of the polypeptide encoded by clone P00258_A10 (SEQ ID NO: 56), a polynucleotide encoding at least about 50 contiguous amino acids from amino acids 1 to 245 of SEQ ID NO: 59, wherein said polynucleotide encodes a polypeptide having at least one biological activity of the polypeptide encoded by clone P00262_C10 (SEQ ID NO: 58), a polynucleotide encoding at least about 50 contiguous amino acids from amino acids 1 to 142 of SEQ ID NO: 63, wherein said polynucleotide encodes a polypeptide having at least one biological activity of the polypeptide encoded by clone P00269_H08 (SEQ ID NO: 62), a polynucleotide encoding at least about 50 contiguous amino acids from amino acids 1 to 49 of SEQ ID NO: 67, wherein said polynucleotide encodes a polypeptide having at least one biological activity of the polypeptide encoded by clone P00628_H02 (SEQ ID NO: 66), a polynucleotide encoding at least about 50 contiguous amino acids from amino acids 1 to 70 of SEQ ID NO: 69, wherein said polynucleotide encodes a polypeptide having at least one biological activity of the polypeptide encoded by clone P00629_C08 (SEQ ID NO: 68), a polynucleotide encoding at least about 50 contiguous amino acids from amino acids 1 to 113 of SEQ ID NO: 72, wherein said polynucleotide encodes a polypeptide having at least one biological activity of the polypeptide encoded by clone P00641_G11 (SEQ ID NO: 71), a polynucleotide encoding at least about 50 contiguous amino acids from amino acids 1 to 114 of SEQ ID NO: 74, wherein said polynucleotide encodes a polypeptide having at least one biological activity of the polypeptide encoded by clone P00648_E12 (SEQ ID NO: 73), a polynucleotide encoding at least about 50 contiguous amino acids

from amino acids 1 to 97 of SEQ ID NO: 76, wherein said polynucleotide encodes a polypeptide having at least one biological activity of the polypeptide encoded by clone P00697_C03 (SEQ ID NO: 75);

5 (f) a polynucleotide encoding at least about 50 contiguous amino acids from amino acids 1 to 23 of SEQ ID NO: 26, wherein said polynucleotide encodes a polypeptide having at least one biological activity of the polypeptide encoded by clone P00219_F06 (SEQ ID NO: 25) or amino acids 1 to 387 of SEQ ID NO: 53, wherein said polynucleotide encodes a polypeptide having at least one biological activity of the polypeptide encoded by clone P00248_B04 (SEQ ID NO: 52);

10 (g) a polynucleotide of SEQ ID NOS:1, 3, 5, 7, 9, 11, 13, 15, 17, 19, 21, 23, 25, 27, 29, 31, 32, 34, 36, 38, 40, 42, 44, 45, 47, 48, 49, 50, 52, 54, 56, 58, 60, 61, 62, 64, 65, 66, 68, 70, 71, 73, and 75;

15 (h) the complement of a polynucleotide of (a) – (g); and

(i) an antisense oligonucleotide capable of hybridizing with, and inhibiting the translation of, the mRNA encoded by a gene encoding a polypeptide of SEQ ID NOS: 2, 4, 6, 8, 10, 12, 14, 16, 18, 20, 22, 24, 26, 28, 30, 33, 35, 37, 39, 41, 43, 46, 51, 53, 55, 57, 59, 63, 67, 69, 72, 74, 76, or another mammalian homologue thereof.

20 2. The polynucleotide of claim 1 encoding a polypeptide comprising amino acids 1 to 233 of SEQ ID NO: 26, amino acids 1 to 387 of SEQ ID NO: 53.

3. The polynucleotide of claim 1 comprising the sequence selected from the group consisting of SEQ ID NOS:1, 3, 5, 7, 9, 11, 13, 15, 17, 19, 21, 23, 25, 27, 29, 31, 32, 34, 36, 38, 40, 42, 44, 45, 47, 48, 49, 50, 52, 54, 56, 58, 60, 61, 62, 64, 65, 66, 68, 70, 71, 73, and 75.

25 4. A vector comprising and capable of expressing a poly- or oligonucleotide of claim 1.

5. A recombinant host cell transformed with nucleic acid comprising a poly- or oligonucleotide of claim 1.

6. A recombinant host cell transformed with the vector of claim 5.

7. A method for producing a polypeptide comprising culturing a recombinant host cell 25 transformed with nucleic acid comprising any of the polynucleotides of claim 1(a) – (g) under conditions such that the polypeptide is expressed, and isolating the polypeptide.

8. A polypeptide comprising:

(a) a polypeptide having at least about 80% identity with amino acids selected from the group consisting of: amino acids 1 to 193 of SEQ ID NO: 4, amino acids 1 to 236 of SEQ ID NO:6, amino acids 1 to 61 of SEQ ID NO: 8, amino acids 1 to 92 of SEQ ID NO:12, amino acids 1 to 86 of SEQ ID NO:14, amino acids 1 to 36 of SEQ ID NO:16, amino acids 1 to 83 of SEQ ID NO:18, amino acids 1 to 82 of SEQ ID NO:20, amino acids 1 to 462 of SEQ ID NO:22, amino acids 1 to 170 of SEQ ID NO:24, amino acids 1 to 30 of SEQ ID NO:28, amino acids 1 to 30 of SEQ ID NO: 35, amino acids 1 to 100 of SEQ ID NO:37, amino acids 1 to 65 of SEQ ID NO:39, amino acids 1 to 46 of SEQ ID NO:43, amino acids 1 to 313 of 35 SEQ ID NO:46, amino acids 1 to 58 of SEQ ID NO:51, amino acids 1 to 58 of SEQ ID NO:55, amino acids 1 to 52 of SEQ ID NO:57, amino acids 1 to 245 of SEQ ID NO:59, amino acids 1 to 142 of SEQ ID NO:63,

amino acids 1 to 49 of SEQ ID NO:67, amino acids 1 to 70 of SEQ ID NO:69, amino acids 1 to 113 of SEQ ID NO:72, and amino acids 1 to 97 of SEQ ID NO:76; or

(b) a polypeptide encoded by nucleic acid hybridizing under stringent conditions with the complement of the coding region selected from the group consisting of: SEQ ID NOS:1, 3, 5, 7, 9, 11, 13, 15, 17, 19, 21, 23, 25, 27, 29, 31, 32, 34, 36, 38, 40, 42, 44, 45, 47, 48, 49, 50, 52, 54, 56, 58, 60, 61, 62, 64, 65, 66, 68, 70, 71, 73, and 75;

(c) the polypeptides of (a) and (b) having at least one biological activity of the polypeptide encoded by clones P00184_D11 (SEQ ID NO:1), P00185_D11(SEQ ID NO:3), P00188_D12 (SEQ ID NO:5), P00188_E01 (SEQ ID NO:7), P00194_G01 (SEQ ID NO:9), P00194_G05 (SEQ ID NO:11), P00194_H10 (SEQ ID NO:13), P00199_D08 (SEQ ID NO:15), P00203_D04 (SEQ ID NO:17), P00203_E06 (SEQ ID NO:19), P00209_F06 (SEQ ID NO:21), P00219_D02 (SEQ ID NO:23), P00219_F06 (SEQ ID NO:25), P00220_H05 (SEQ ID NO:27), P00222_G03 (SEQ ID NO:29), P00225_C01 (SEQ ID NO:32), P00227_D11 (SEQ ID NO:34), P00228_F03 (SEQ ID NO:36), P00233_H08 (SEQ ID NO:38), P00235_G08 (SEQ ID NO:40), P00239_C11 (SEQ ID NO:42), P00240_E05 (SEQ ID NO:45), P00247_A04 (SEQ ID NO:50), P00248_B04 (SEQ ID NO:52), P00249_F09 (SEQ ID NO:54), P00258_A10 (SEQ ID NO:56), P00262_C10 (SEQ ID NO:58), P00269_H08 (SEQ ID NO:62), P00628_H02 (SEQ ID NO:66), P00629_C08 (SEQ ID NO:68), P00641_G11 (SEQ ID NO:71), P00648_E12 (SEQ ID NO:73), P00697_C03 (SEQ ID NO:75).

9. A composition comprising a polypeptide which comprises:

(a) a polypeptide having at least about 80% identity with amino acids selected from the group consisting of: amino acids 1 to 193 of SEQ ID NO: 4, amino acids 1 to 236 of SEQ ID NO:6, amino acids 1 to 61 of SEQ ID NO: 8, amino acids 1 to 92 of SEQ ID NO:12, amino acids 1 to 86 of SEQ ID NO:14, amino acids 1 to 36 of SEQ ID NO:16, amino acids 1 to 83 of SEQ ID NO:18, amino acids 1 to 82 of SEQ ID NO:20, amino acids 1 to 462 of SEQ ID NO:22, amino acids 1 to 170 of SEQ ID NO:24, amino acids 1 to 30 of SEQ ID NO:28, amino acids 1 to 30 of SEQ ID NO: 35, amino acids 1 to 100 of SEQ ID NO:37, amino acids 1 to 65 of SEQ ID NO:39, amino acids 1 to 46 of SEQ ID NO:43, amino acids 1 to 313 of SEQ ID NO:46, amino acids 1 to 58 of SEQ ID NO:51, amino acids 1 to 58 of SEQ ID NO:55, amino acids 1 to 52 of SEQ ID NO:57, amino acids 1 to 245 of SEQ ID NO:59, amino acids 1 to 142 of SEQ ID NO:63, amino acids 1 to 49 of SEQ ID NO:67, amino acids 1 to 70 of SEQ ID NO:69, amino acids 1 to 113 of SEQ ID NO:72, and amino acids 1 to 97 of SEQ ID NO:76; or

(b) a polypeptide encoded by nucleic acid hybridizing under stringent conditions with the complement of the coding region selected from the group consisting of: SEQ ID NOS:1, 3, 5, 7, 9, 11, 13, 15, 17, 19, 21, 23, 25, 27, 29, 31, 32, 34, 36, 38, 40, 42, 44, 45, 47, 48, 49, 50, 52, 54, 56, 58, 60, 61, 62, 64, 65, 66, 68, 70, 71, 73, and 75; wherein the polypeptides of (a) and (b) have at least one biological activity of the polypeptide respectively encoded by clones P00184_D11 (SEQ ID NO:1), P00185_D11(SEQ ID NO:3), P00188_D12 (SEQ ID NO:5), P00188_E01 (SEQ ID NO:7), P00194_G01 (SEQ ID NO:9), P00194_G05 (SEQ ID NO:11), P00194_H10 (SEQ ID NO:13), P00199_D08 (SEQ ID

NO:15), P00203_D04 (SEQ ID NO:17), P00203_E06 (SEQ ID NO:19), P00209_F06 (SEQ ID NO:21), P00219_D02 (SEQ ID NO:23), P00219_F06 (SEQ ID NO:25), P00220_H05 (SEQ ID NO:27), P00222_G03 (SEQ ID NO:29), P00225_C01 (SEQ ID NO:32), P00227_D11 (SEQ ID NO:34), P00228_F03 (SEQ ID NO:36), P00233_H08 (SEQ ID NO:38), P00235_G08 (SEQ ID NO:40),
5 P00239_C11 (SEQ ID NO:42), P00240_E05 (SEQ ID NO:45), P00247_A04 (SEQ ID NO:50), P00248_B04 (SEQ ID NO:52), P00249_F09 (SEQ ID NO:54), P00258_A10 (SEQ ID NO:56), P00262_C10 (SEQ ID NO:58), P00269_H08 (SEQ ID NO:62), P00628_H02 (SEQ ID NO:66), P00629_C08 (SEQ ID NO:68), P00641_G11 (SEQ ID NO:71), P00648_E12 (SEQ ID NO:73), and P00697_C03 (SEQ ID NO:75), in admixture with a carrier.

10 10. The composition of claim 9 which is a pharmaceutical composition comprising an effective amount of said polypeptide in admixture with a pharmaceutically acceptable carrier.

11. An antibody specifically binding a polypeptide of claim 8.

12. A composition comprising an antibody of claim 11 in admixture with a carrier.

15 13. The composition of claim 9 which is a pharmaceutical composition comprising an effective amount of said antibody in admixture with a pharmaceutically acceptable carrier.

14. A composition comprising an antagonist or an agonist of a polypeptide of claim 8.

15. The composition of claim 11 which is a pharmaceutical composition comprising an effective amount of said antagonist or said agonist in combination with a pharmaceutically acceptable carrier.

20 16. A method for the treatment of a cardiac, renal or inflammatory disease, comprising administering to a patient in need an effective amount of a polypeptide of claim 8, or an antagonist or agonist thereof.

17. A method for the treatment of a cardiac, renal or inflammatory disease, comprising administering to a patient in need an effective amount of an antibody specifically binding to a polypeptide of the present invention.

25 18. A method for screening a subject for a cardiac, renal or inflammatory disease characterized by the differential expression of the polypeptide selected from the group consisting of: SEQ ID NOS: 2, 4, 6, 8, 10, 12, 14, 16, 18, 20, 22, 24, 26, 28, 30, 33, 35, 37, 39, 41, 43, 46, 51, 53, 55, 57, 59, 63, 67, 69, 72, 74, and 76, or an endogenous homologue thereof, comprising the steps of:

measuring the expression in the subject of said polypeptide or said endogenous homologue; and

30 determining the relative expression of said polypeptide or said endogenous homologue in the subject compared to its expression in normal subjects, or compared to its expression in the same subject at an earlier stage of development of the cardiac, renal or inflammatory disease.

35 19. The method of claim 15 wherein said subject is human and said endogenous homologue is a human homologue of the rat protein selected from the group consisting of: SEQ ID NOS: 2, 4, 6, 8, 10, 12, 14, 16, 18, 20, 22, 24, 26, 28, 30, 33, 35, 37, 39, 41, 43, 46, 51, 53, 55, 57, 59, 63, 67, 69, 72, 74, and 76.

20. An array comprising one or more oligonucleotides complementary to reference RNA or

DNA encoding a protein selected from the group consisting of: SEQ ID NOS: 2, 4, 6, 8, 10, 12, 14, 16, 18, 20, 22, 24, 26, 28, 30, 33, 35, 37, 39, 41, 43, 46, 51, 53, 55, 57, 59, 63, 67, 69, 72, 74, and 76, or another mammalian (e.g. human) homologue thereof, where the reference DNA or RNA sequences are obtained from both a biological sample from a normal subject and a biological sample from a subject exhibiting a
5 cardiac, renal, or inflammatory disease, or from biological samples taken at different stages of a cardiac, renal, or inflammatory disease.

21. A method for detecting cardiac, kidney, or inflammatory disease in a human test patient comprising the steps of:

providing an array of oligonucleotides at known locations on a substrate, which array comprises
10 oligonucleotides complementary to reference DNA or RNA sequences encoding a human homologue of the protein selected from the group consisting of: SEQ ID NOS: 2, 4, 6, 8, 10, 12, 14, 16, 18, 20, 22, 24, 26, 28, 30, 33, 35, 37, 39, 41, 43, 46, 51, 53, 55, 57, 59, 63, 67, 69, 72, 74, and 76 where the reference DNA or RNA sequences are obtained from both a biological sample from a normal patient and a biological sample from a patient potentially exhibiting cardiac, renal, or inflammatory disease, or from a test patient
15 exhibiting cardiac, renal, or inflammatory disease, taken at different stages of such disease;

exposing the array, under hybridization conditions, to a first sample of cDNA probes constructed from mRNA obtained from a biological sample from a corresponding biological sample of a normal patient or from a test patient at a certain stage of the disease;

exposing the array, under hybridization conditions, to a second sample of cDNA probes
20 constructed from mRNA obtained from a biological sample obtained from the test;

quantifying any hybridization between the first sample of cDNA probes and the second sample of cDNA probes with the oligonucleotide probes on the array; and

determining the relative expression of genes encoding the human homologue of a protein selected
25 from the group consisting of: SEQ ID NOS: 2, 4, 6, 8, 10, 12, 14, 16, 18, 20, 22, 24, 26, 28, 30, 33, 35, 37, 39, 41, 43, 46, 51, 53, 55, 57, 59, 63, 67, 69, 72, 74, and 76 in the biological samples from the normal patient and the test patient, or in the biological samples taken from the test patient at different stages of the disease.

22. A diagnostic kit for the detection of a cardiac, kidney or inflammatory disease comprising an array of claim 20.

30 23. The diagnostic kit of claim 22 further comprising at least one of the following components:

- (a) an oligonucleotide probe;
- (b) a PCR reagent;
- (c) a detectable label;
- 35 (d) a biological sample taken from a human subject; and

(e) an antibody to a polypeptide of any one of the sequences selected from the group consisting of: SEQ ID NOS: 2, 4, 6, 8, 10, 12, 14, 16, 18, 20, 22, 24, 26, 28, 30, 33, 35, 37, 39, 41, 43, 46, 51, 53, 55, 57, 59, 63, 67, 69, 72, 74, 76, and a further mammalian homologue thereof.

24. The diagnostic kit of claim 22 wherein said biological sample is from blood or a tissue.

5 25. The diagnostic kit of claim 21 wherein said tissue is a cardiac tissue.

26. The diagnostic kit of claim 22 wherein said cardiac tissue is a left ventricular tissue.

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FIGURE 38

5	tctagcgaac cccttcgtag aactaggagc cagtgttgc acggcggt ggctggatac cccactgcat gctgcagcaa ggcagtccag tgtggaggtc atcaatctgc tcactgagta tggggctaactca ctgaaactca gaaaactcgca gggcaaaagt gctcttgagc tcgctgctcc caaaagtagt gtggagcagg cactcctgct ccatgaaggt ccacctgctc tttctcagct ctgccgcttg tgtgtccgga agtgcttggg ccgcac atg tca tca agc cat cta	60 120 180 240 294
10	Met Ser Ser Ser His Leu 1 5	
15	cgc act agg tct gcc aga acc cct gga aaa att cct ctt ata cca ata Arg Thr Arg Ser Ala Arg Thr Pro Gly Lys Ile Pro Leu Ile Pro Ile 10 15 20	342
20	gtt gga aac atg ttg cct gct gta gga cac tta ata tac aca ttc agt Val Gly Asn Met Leu Pro Ala Val Gly His Leu Ile Tyr Thr Phe Ser 25 30 35	390
25	ggc tta acc cac tat cct aaa aat ctg ctt acc taa ttagaataaa Gly Leu Thr His Tyr Pro Lys Asn Leu Leu Thr * 40 45	436
30	gccttcataa atccaaatac ttgcgttcaa caaactcctg gtttaggttaa tggntgcca gagataacca gaaaccttcc aagttttaa ctcttggtaa tttaaaatca aactgaaata gatggaaaat aataatctat ttttggataa ttcaaggacc ctgcgtatc tggggctgg gtccgcattt tgnatactgg atagacacac acacaggtag gatanggtaa atnaactact taaagaatgg cctgggattt aagtccctcca gatattttt aggtngnggt ttccctaaaaat 35 40 45 50 55 60 65 70 75 80 85 90 95 100 105 110 115 120 125 130 135 140 145 150 155 160 165 170 175 180 185 190 195 200 205 210 215 220 225 230 235 240 245 250 255 260 265 270 275 280 285 290 295 300 305 310 315 320 325 330 335 340 345 350 355 360 365 370 375 380 385 390 395 400 405 410 415 420 425 430 435 440 445 450 455 460 465 470 475 480 485 490 495 500 505 510 515 520 525 530 535 540 545 550 555 560 565 570 575 580 585 590 595 600 605 610 615 620 625 630 635 640 645 650 655 660 665 670 675 680 685 690 695 700 705 710 715 720 725 730 735 740 745 750 755 760 765 770 775 780 785 790 795 800 805 810 815 820 825 830 835 840 845 850	496 556 616 676 736 782

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Sl/Sq

